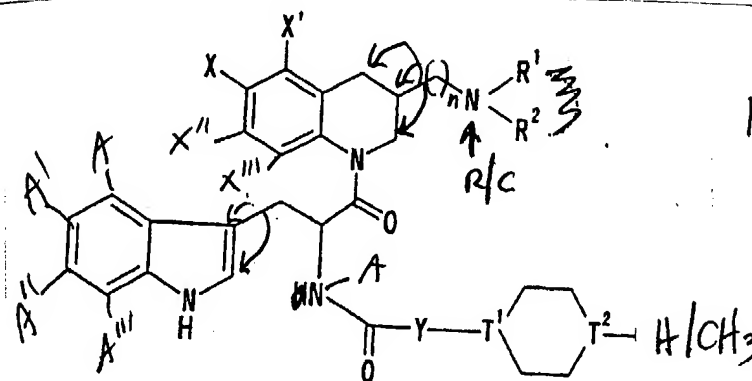


SEARCH REQUEST FORM 131327

Requestor's Name: BERCH Serial Number: 10/089251
Date: 8/31/04 Phone: 571-272-0663 Art Unit: 1624
Office Rem 5C01 Mailbox 5C18 MEJ

Search Topic:

Please write a detailed statement of search topic. Describe specifically as possible the subject matter to be searched. Define any terms that may have a special meaning. Give examples or relevant citations; authors, keywords, etc., if known. For sequences, please attach a copy of the sequence. You may include a copy of the broadest and/or most relevant claim(s).



all $X = H/Cl/F/CH_3$ but exclude the situations where all are H

$R^1, R^2 = H/C$

Y = bond a chain of 1-6 atoms $O/S/N/C$

$T^1, T^2 = C/N$

12:10
16:48 - 58
8

323
323.31
369.54
69285

1648

Buch
10/089951

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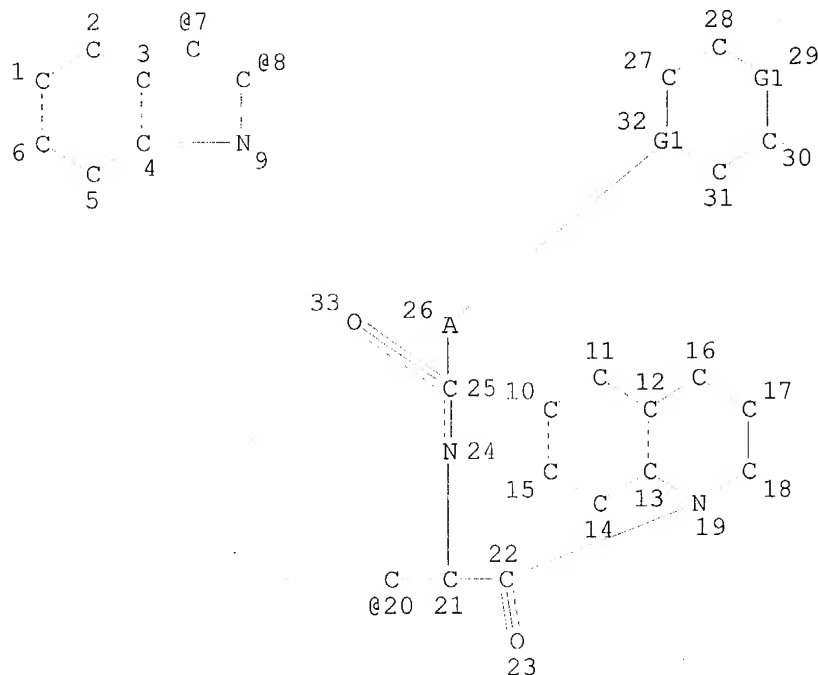
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FILE 'REGISTRY' ENTERED AT 16:48:27 ON 31 AUG 2004

L1 STR
L2 0 S L1
L3 1 S L1 FUL

=> d l3 que stat;d ide cbib abs

L1 STR



VAR G1=C/N
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DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 33

STEREO ATTRIBUTES: NONE
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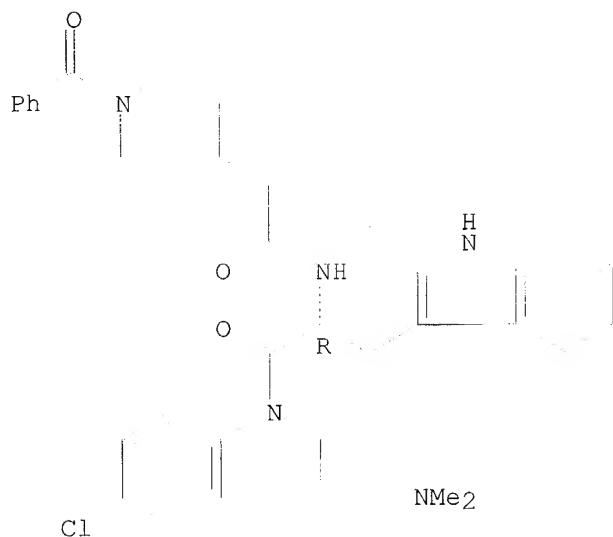
1 ANSWERS

L3 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2004 ACS on STN
RN 333952-73-1 REGISTRY
CN 4-Piperidineacetamide, 1-benzoyl-N-[(1R)-2-[6-chloro-3-
[(dimethylamino)methyl]-3,4-dihydro-1(2H)-quinolinyl]-1-(1H-indol-3-
ylmethyl)-2-oxoethyl]- (9CI) (CA INDEX NAME)

Searched by: Mary Hale 571-272-2507 REM 1D86

FS STEREOSEARCH
 MF C37 H42 Cl N5 O3
 SR CA
 LC STN Files: CA, CAPLUS, TOXCENTER
 DT.CA Caplus document type: Patent
 RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.

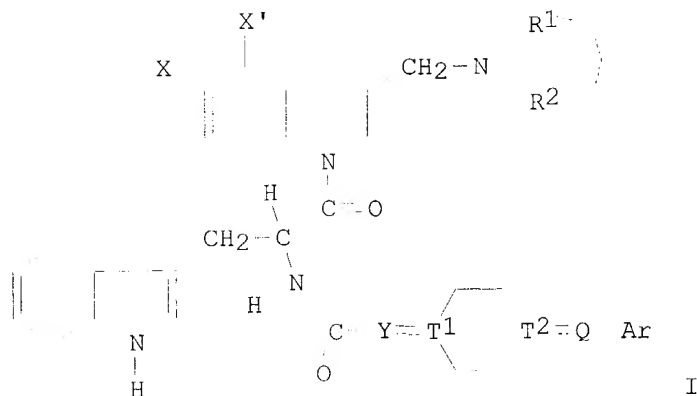


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
 1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

REFERENCE 1: 134:295840 Preparation of indolylpropanoyltetrahydroquinoline derivatives which inhibit binding of somatostatin receptors. Kato, Kaneyoshi; Terauchi, Jun; Suzuki, Nobuhiro; Takekawa, Shiro (Tadeca Chemical Industries, Ltd., Japan). PCT Int. Appl. WO 2001025228 A1 20010412, 220 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MA, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (Japanese). CODEN: PIXXD2. APPLICATION: WO 2000-JP6937 20001005. PRIORITY: JP 1999-286939 19991007; JP 2000-215837 20000711.

GI



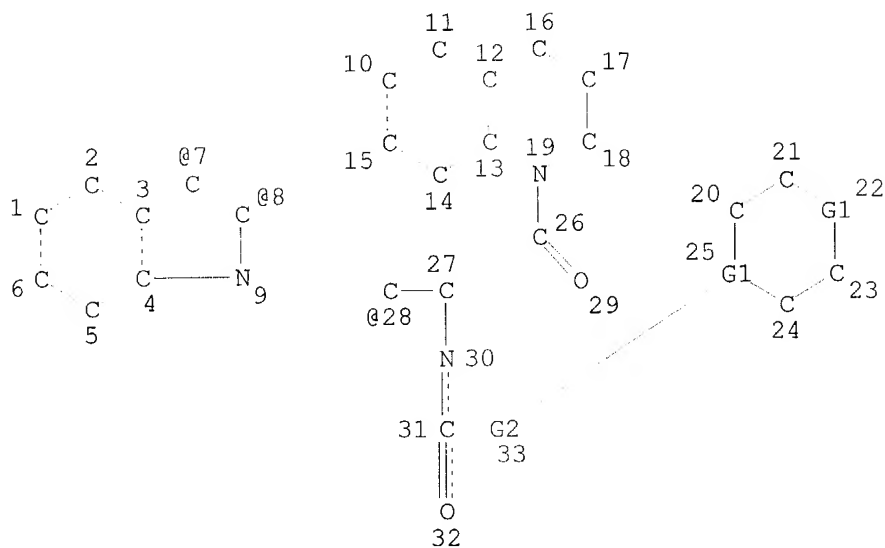
AB The title compds. I [X and X' are the same or different and each represents hydrogen, fluorine, etc., provided that at least one of X and X' represents fluorine, chlorine, etc.; R1 and R2 represents each hydrogen or optionally substituted C1-6 alkyl, or R1 and R2 form together with the nitrogen atom adjacent thereto an optionally substituted nitrogen-containing heterocycle; Y and Q are the same or different and each represents a bond or a spacer having 1 to 6 atoms in the main chain; the dotted line represents a single or double bond; T1 and T2 represent each C(R9) (wherein R9 represents hydrogen, hydroxy, etc.), N, etc.; and Ar represents an optionally substituted aromatic group, hydrogen, etc.; a provision is given] are prepared In an in vitro test for inhibition of binding to the somatostatin receptor type 2, several compds. of this invention showed IC50 of 0.6 to 2 nM. Formulations are given.

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L7 0 L6 NOT L3

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	ENTRY	SESSION
FULL ESTIMATED COST	323.10	323.31
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE	TOTAL
	ENTRY	SESSION
CA SUBSCRIBER PRICE	-0.66	-0.66

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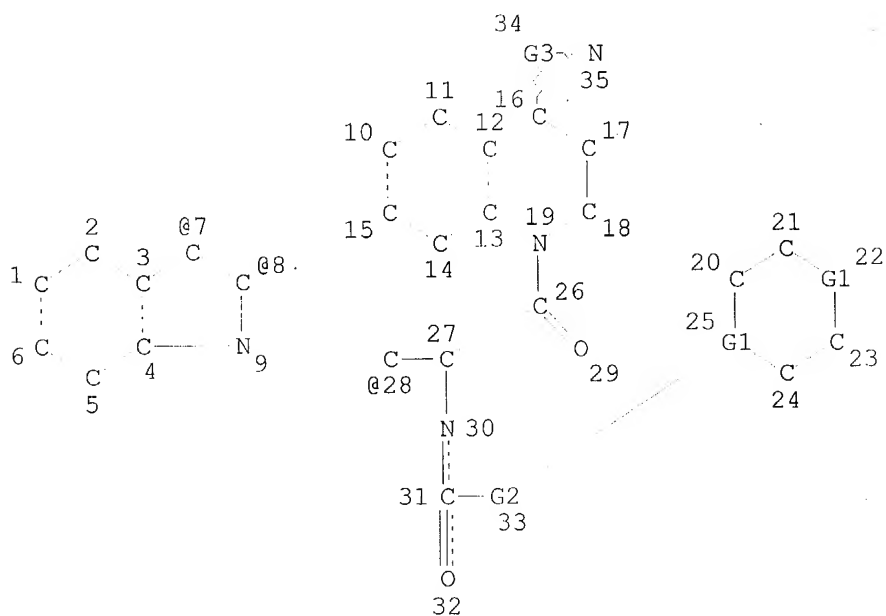
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REP G2=(0-6) A
VPA 28-7/8 U
NODE ATTRIBUTES:
DEFAULT MLEVEL IS ATOM
DEFAULT ECLEVEL IS LIMITED
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GRAPH ATTRIBUTES:
RING(S) ARE ISOLATED OR EMBEDDED
NUMBER OF NODES IS 33
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STEREO ATTRIBUTES: NONE
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L4 STR
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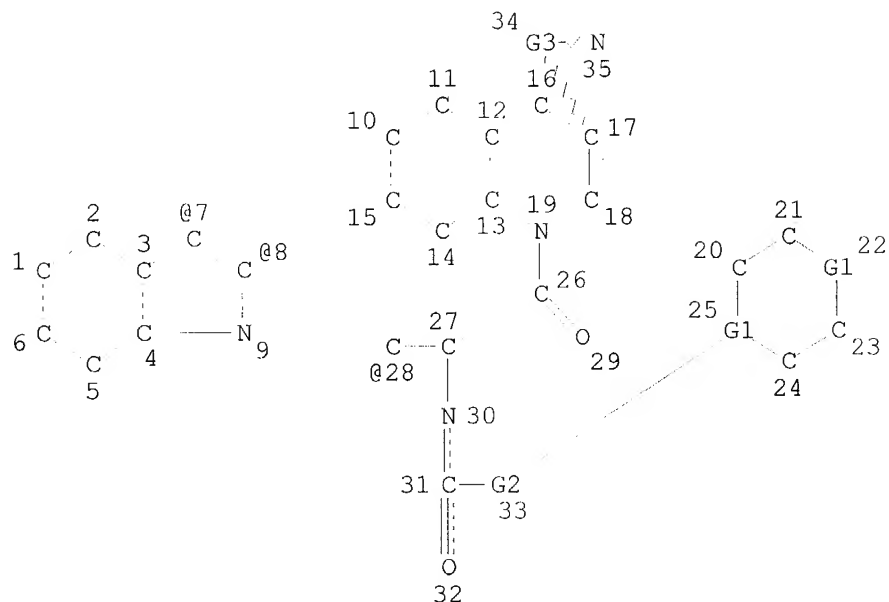
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Searched by: Mary Hale 571-272-2507 REM 1D86

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 REP G3=(1-3) C
 VPA 28-7/8 U
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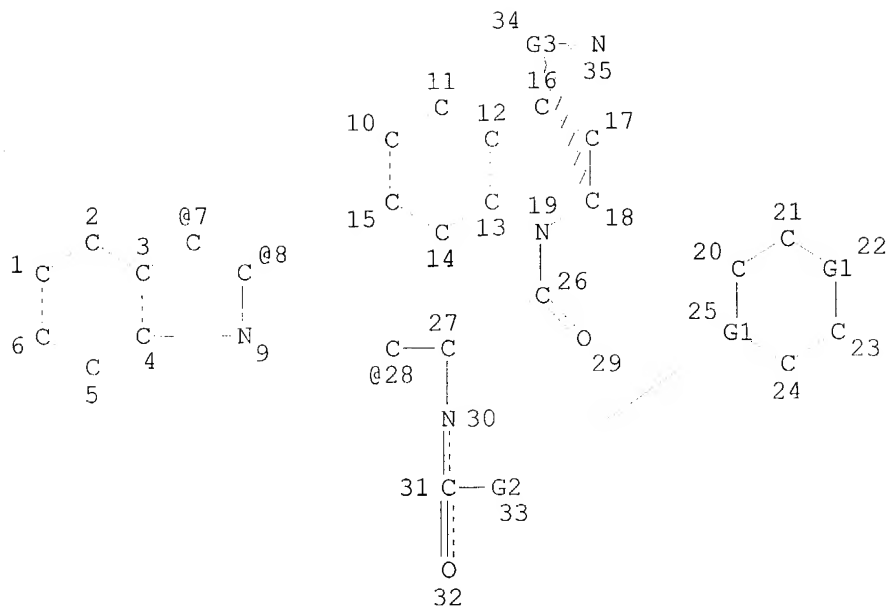
STEREO ATTRIBUTES: NONE
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VAR G1=C/N
 REP G2=(0-6) A
 REP G3=(1-3) C
 VPA 28-7/8 U
 NODE ATTRIBUTES:
 NSPEC IS RC AT 35
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 35

STEREO ATTRIBUTES: NONE
 L6 STR



VAR G1=C/N
 REP G2=(0-6) A
 REP G3=(1-3) C
 VPA 28-7/8 U
 NODE ATTRIBUTES:
 NSPEC IS RC AT 35
 DEFAULT MLEVEL IS ATOM
 DEFAULT ECLEVEL IS LIMITED

GRAPH ATTRIBUTES:
 RING(S) ARE ISOLATED OR EMBEDDED
 NUMBER OF NODES IS 35

STEREO ATTRIBUTES: NONE
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100.0% PROCESSED 303 ITERATIONS 298 ANSWERS
 SEARCH TIME: 00.00.01

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	196.69	196.90

FILE 'HCAPLUS' ENTERED AT 17:06:39 ON 31 AUG 2004
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FILE COVERS 1907 - 31 Aug 2004 VOL 141 ISS 10
FILE LAST UPDATED: 30 Aug 2004 (20040830/ED)

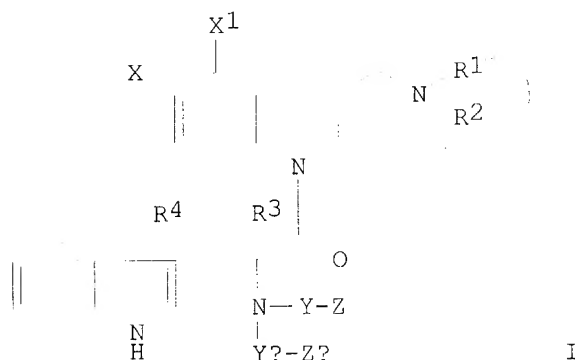
This file contains CAS Registry Numbers for easy and accurate substance identification.

L9 3 L8

=> d 1-3 cbib abs

L9 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
2003:396877 Document No. 138:401769 Preparation of [1-[3-(indol-3-yl)propanoyl]-1,2,3,4-tetrahydroquinolin-3-ylmethyl]amine derivatives as somatostatin receptor binding inhibitors. Abe, Hidenori; Kasai, Shizuo; Takekawa, Shiro; Watanabe, Masanori (Takeda Chemical Industries, Ltd., Japan). PCT Int. Appl. WO 2003042204 A1 20030522, 191 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR. (Japanese). CODEN: PIXXD2. APPLICATION: WO 2002-JP10800 20021017. PRIORITY: JP 2001-322897 20011019.

GI



AB The title compds. represented by the formula (I) (wherein X and X1 are the same or different and each represents H, halo, or (un)substituted NH₂; R1 and R2 are the same or different and each represents H or (un)substituted C1-6 alkyl; or NR1R2 forms (un)substituted N-containing heterocyclic ring; R3 represents an each optionally substituted hydrocarbon group or heterocyclyl; R4 represents H or an each optionally substituted hydrocarbon group or heterocyclyl; Y and Ya are the same or different and each represents a bond or a spacer having a C1-8 main chain; and Z and Za are the same or different and each represents H, halo, or (un)substituted

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cyclic group), salts of the compds., or prodrugs of either are prepared. They have inhibitory activity against somatostatin receptor, in particular somatostatin receptor subtype 2 binding and are agonists of somatostatin receptor and effective in the prevention of and treatment for diseases in which somatostatin participates, in particular diabetes or diabetes complications. Thus, a solution of 2.6 g (2RS,3SR)-2-[[[9H-fluoren-9-ylmethoxy)carbonyl]amino]-3-(1H-indol-3-yl)butanoic acid and 0.06 mL DMF in 60 mL THF was treated dropwise with a solution of 0.63 mL oxalyl chloride in 5 mL THF at 0°, stirred at 0° for 30 min, concentrated, treated with 30 mL THF, and reconcd., dissolved in 30 mL THF, added dropwise at 0° to a solution of 1-[(3S)-6-chloro-1,2,3,4-tetrahydroquinolin-3-yl]-N,N-dimethylmethanamine 0.90 g, tetrabutylammonium hydrogen sulfate 0.04 g, and NaOH powder 0.34 g, stirred at 0° for 30 min to give, after workup and silica gel chromatog., a yellow amorphous solid which was stirred with 0.2 mL piperidine in 20 mL methanol at room temperature for 16 h

to

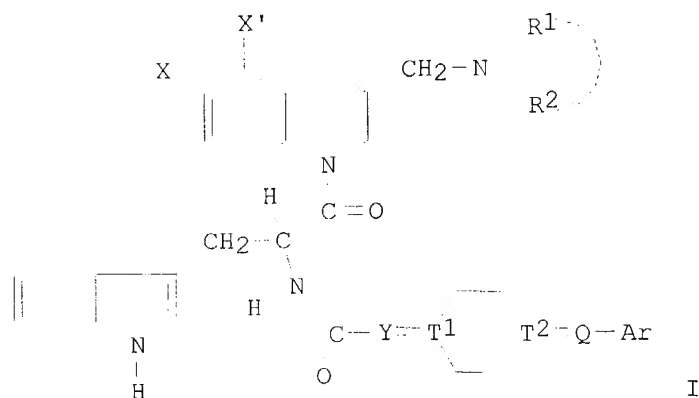
give, after alumina chromatog., 49% (2RS,3SR)-1-[(3R)-6-chloro-3-[(dimethylamino)methyl]-3,4-dihydro-1(2H)-quinolinyl]-3-(1H-indol-3-yl)-1-oxo-2-butanamine (II; R = H). WSC (0.10 g) was added to a solution of II 0.20, 1-[(1-methyl-1H-indol-2-yl)carbonyl]-4-piperidinecarboxylic acid 0.15 g, and HOBt 0.08 g in 10 mL MeCN, stirred at room temperature for 16 h to give, after workup and silica gel chromatog., 64% II (R = Q). II (R = Q1) in vitro inhibited the binding of 125I-somatostatin-14 to human somatostatin receptor protein subtype 2, 3, and 5 with showed IC50 of 0.05, 3, and 10, resp. A tablet formulation containing II (R = H) was described.

L9 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN

2001:265411 Document No. 134:295840 Preparation of

indolylpropanoyltetrahydroquinoline derivatives which inhibit binding of somatostatin receptors. Kato, Kaneyoshi; Terauchi, Jun; Suzuki, Nobuhiro; Takekawa, Shiro (Tadaka Chemical Industries, Ltd., Japan). PCT Int. Appl. WO 2001025228 A1 20010412, 220 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CN, CR, CU, CZ, DM, DZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MA, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, TJ, TM, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (Japanese). CODEN: PIXXD2. APPLICATION: WO 2000-JP6937 20001005. PRIORITY: JP 1999-286939 19991007; JP 2000-215837 20000711.

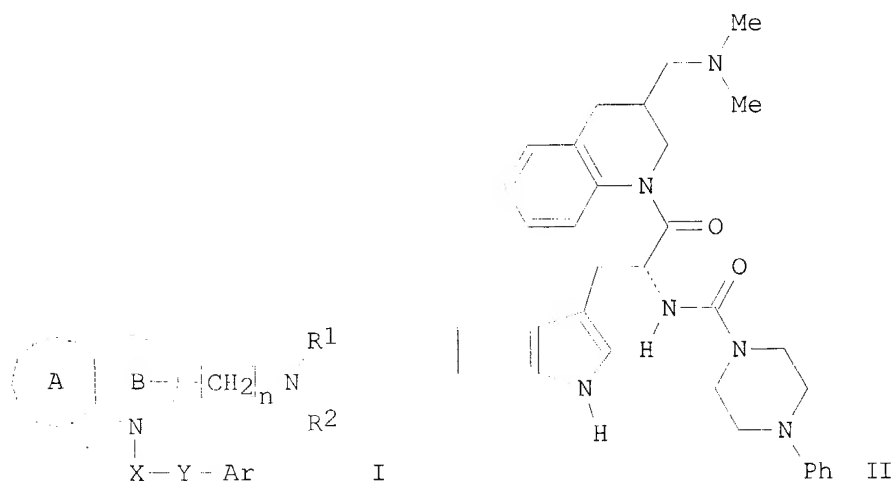
GI



AB The title compds. I [X and X' are the same or different and each represents hydrogen, fluorine, etc., provided that at least one of X and X' represents fluorine, chlorine, etc.; R1 and R2 represents each hydrogen or optionally substituted C1-6 alkyl, or R1 and R2 form together with the nitrogen atom adjacent thereto an optionally substituted nitrogen-containing heterocycle; Y and Q are the same or different and each represents a bond or a spacer having 1 to 6 atoms in the main chain; the dotted line represents a single or double bond; T1 and T2 represent each C(R9) (wherein R9 represents hydrogen, hydroxy, etc.), N, etc.; and Ar represents an optionally substituted aromatic group, hydrogen, etc.; a provision is given] are prepared In an in vitro test for inhibition of binding to the somatostatin receptor type 2, several compds. of this invention showed IC50 of 0.6 to 2 nM. Formulations are given.

L9 ANSWER 3 OF 3. HCAPLUS COPYRIGHT 2004 ACS on STN
 1999:672759 Document No. 131:286420 Preparation of amine compounds as somatostatin receptor antagonists or agonists. Suzuki, Nobuhiro; Kato, Kaneyoshi; Takekawa, Shiro; Terauchi, Jun; Endo, Satoshi (Takeda Chemical Industries, Ltd., Japan). PCT Int. Appl. WO 9952875 A1 19991021, 257 pp. DESIGNATED STATES: W: AE, AL, AM, AU, AZ, BA, BB, BG, BR, BY, CA, CN, CU, CZ, EE, GD, GE, HR, HU, ID, IL, IN, IS, JP, KG, KR, KZ, LC, LK, LR, LT, LV, MD, MG, MK, MN, MX, NO, NZ, PL, RO, RU, SG, SI, SK, SL, TJ, TM, TR, TT, UA, US, UZ, VN, YU, ZA, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 1999-JP1871 19990408. PRIORITY: JP 1998-96422 19980408; JP 1998-345328 19981204.

GI



AB The title compds. [I; Ar = (un)substituted aromatic; X = CH₂, S, SO, SO₂, CO; Y = a spacer having a main chain of 2-5 atoms; n = 1-5; R1, R2 = H, lower alkyl; NR1R2 = (un)substituted nitrogen-containing heterocyclic ring; R1 or R2 together with -(CH₂)_n-N= form, bonded to a component atom of Ring B, a spiro-ring which may be substituted; Ring A = (un)substituted aromatic; Ring B = (un)substituted 4-7 membered nitrogen-containing non-aromatic ring, with a proviso that X = S, SO, SO₂, CO when Ring A has as a substituent a group -NHCOR11 (wherein R11 = alkyl, alkoxyalkyl, alkylthioalkyl, etc.) or a group NHR12 (R12 = alkyl, cycloalkyl, cycloalkylalkyl, etc.)] or their

salts which have an excellent somatostatin receptor binding inhibition action and are useful for preventing or treating glaucoma, acromegaly, diabetes, diabetic complications or tumor, and as analgesics, were prepared. Thus, treatment of 1-[2-(R)-amino-3-(indol-3-yl)propanoyl]-3-(R,S)-(N,N-dimethylamino)methyl-1,2,3,4-tetrahydroquinoline (preparation described) with N,N'-disuccinimidyl carbonate and N-ethyl-diisopropylamine in THF followed by the addition of solution of 1-phenylpiperazine and N-ethyl-diisopropylamine

in

THF afforded II which showed IC₅₀ of 0.009 μ M and 0.0008 μ M against SSSTR2 and SSSTR3 binding, resp.

=> fil caol;s 19

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
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DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	-2.10	-2.10

FILE 'CAOLD' ENTERED AT 17:06:52 ON 31 AUG 2004
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FILE COVERS 1907-1966
 FILE LAST UPDATED: 01 May 1997 (19970501/UP)

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L10 0 L8

=> fil hcapl;select rn 18

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
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CA SUBSCRIBER PRICE	0.00	-2.10

FILE 'HCAPLUS' ENTERED AT 17:07:09 ON 31 AUG 2004
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Searched by: Mary Hale 571-272-2507 REM 1D86

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FILE COVERS 1907 - 31 Aug 2004 VOL 141 ISS 10
FILE LAST UPDATED: 30 Aug 2004 (20040830/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

NO ANSWERS SELECTED. THE ANSWER SET WAS CREATED IN FILE 'REGISTRY'.
USE THE FILE COMMAND TO CHANGE TO THE CORRECT FILE.
An answer set can be processed to create terms only in the same file in which it was created.

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ENTER ANSWER NUMBER OR RANGE (1-):1-3
E1 THROUGH E829 ASSIGNED

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FULL ESTIMATED COST	2.60	209.93
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FILE 'REGISTRY' ENTERED AT 17:07:44 ON 31 AUG 2004
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STRUCTURE FILE UPDATES: 30 AUG 2004 HIGHEST RN 736108-36-4
DICTIONARY FILE UPDATES: 30 AUG 2004 HIGHEST RN 736108-36-4

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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at:
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DR	107022-57-1		
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DR	105459-20-9		
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DR	31074-85-8		
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790	RN	528-75-6	REGISTRY
791	RN	506-59-2	REGISTRY
792	RN	501-53-1	REGISTRY
DR	94274-21-2		
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794	RN	383-63-1	REGISTRY

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804	RN	108-24-7	REGISTRY
805	RN	108-05-4	REGISTRY
DR	172702-77-1,	61891-42-7, 85306-26-9, 82041-23-4, 220713-36-0	
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808	RN	106-48-9	REGISTRY
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DR	170016-12-3		
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DR	719304-66-2		
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813	RN	100-52-7	REGISTRY
814	RN	100-39-0	REGISTRY
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817	RN	98-09-9	REGISTRY
DR	114415-79-1		
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DR	61840-56-0, 110617-59-9		
820	RN	92-54-6	REGISTRY
DR	123317-04-4		
821	RN	90-02-8	REGISTRY
822	RN	79-43-6	REGISTRY
DR	42428-47-7		
823	RN	79-03-8	REGISTRY
824	RN	75-36-5	REGISTRY
825	RN	75-31-0	REGISTRY
DR	85404-24-6		
826	RN	75-07-0	REGISTRY
827	RN	74-11-3	REGISTRY
828	RN	70-11-1	REGISTRY
829	RN	50-78-2	REGISTRY
DR	11126-35-5, 11126-37-7, 98201-60-6, 2349-94-2, 26914-13-6		

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COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
159.61	369.54

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
0.00	-2.10

CA SUBSCRIBER PRICE

STN INTERNATIONAL LOGOFF AT 17:10:51 ON 31 AUG 2004